

CERTIFICATE OF MAILING

I hereby certify that this correspondence is being deposited with the United States Postal Service as first class mail in an envelope addressed to: Assistant Commissioner for Patents, Washington, D.C. 20231, on the date appearing below.  
ELI LILLY AND COMPANY

By Shonda M. Swab:m

Date February 19, 1999

FEB 24 1999

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

PATENT

U.S. Patent No. : 5,250,542  
Issued: : October 5, 1993  
On an Application of : Buddy E. Cantrell, et al.  
Serial No. : 07/916,783  
Filed : July 17, 1992  
Entitled : PERIPHERALLY SELECTIVE PIPERIDINE  
CARBOXYLTE OPIOID ANTAGONISTS  
Docket No. : x-8244a

APPROVED

OCT 26 1999

FOR THE COMMISSIONER OF PATENT & TRADEMARK OFFICE

REQUEST FOR CERTIFICATE OF CORRECTION

UNDER 37 C.F.R. 1.322

Assistant Commissioner for Patents  
Washington, D. C. 20231  
Sir:

The patentees of the above-identified patent respectfully request that you issue a Certificate of Correction to correct errors which occurred during the printing of the patent. Attached are two copies of Form PTO 1050 on which the errors are specified.

Respectfully submitted,

Nelsen L. Lentz  
Nelsen L. Lentz  
Attorney for Patentees  
Registration No. 38,537  
Phone: 317-276-1207

Eli Lilly and Company  
Patent Division/NLL  
Lilly Corporate Center  
Indianapolis, Indiana 46285

2-19-99

UNITED STATES PATENT AND TRADEMARK OFFICE  
CERTIFICATE OF CORRECTION

PATENT NO. : 5,250,542  
DATED : October 5, 1993  
INVENTOR(S) : Buddy E. Cantrell, et al.

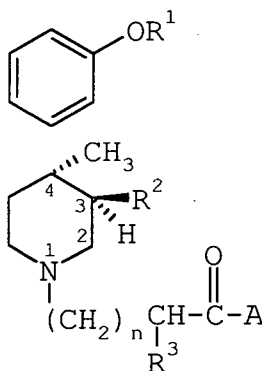
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It is certified that an error appears in the above-identified patent and that said Letters Patent is hereby corrected as shown below:

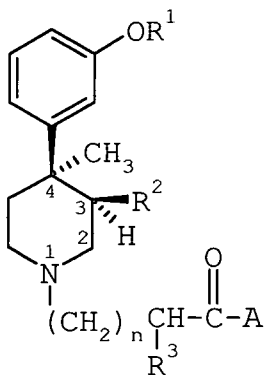
Column 2, line 62, delete the term "CY" and replace it with  $--(CH_2)_n\overset{\overset{O}{\parallel}}{C}Y--$ .

Column 3, line 17, delete the term "C<sub>10</sub>-C<sub>10</sub>" and replace it with  $--(C_1-C_{10})--$ .

Column 5, lines 1-12, delete the structure " and



replace it with --



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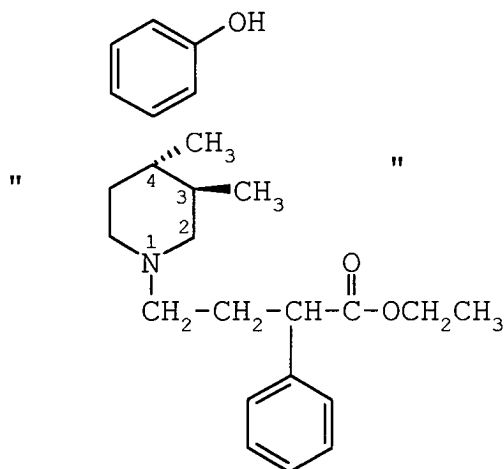
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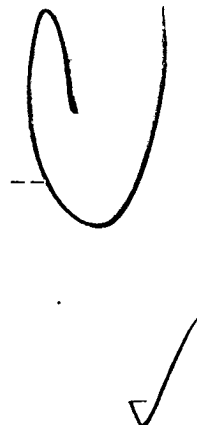
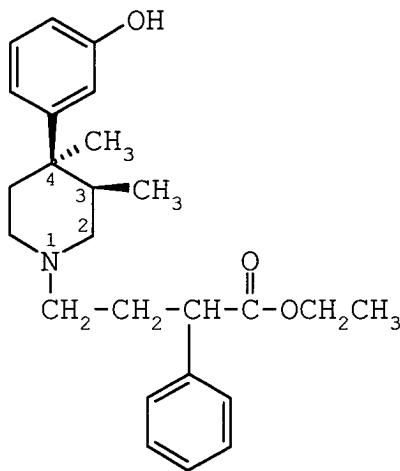
It is certified that an error appears in the above-identified patent and that said Letters Patent is hereby corrected as shown below:

Column 7, lines 12-27, delete the structure



--

and replace it with



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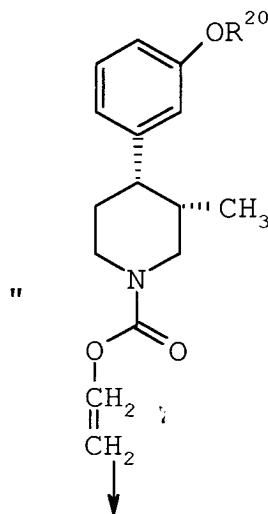
It is certified that an error appears in the above-identified patent and that said Letters Patent is hereby corrected as shown below:

Column 8, line 22, delete the term "pheny)" and replace it with --phenyl)--.

Column 13, line 54, delete the term "R<sup>8</sup>" and replace it with --R<sup>3</sup>--.

Column 13, line 66, delete the term "e-methylene" and replace it with --α-methylene--.

Column 14, lines 1-15, delete the structure



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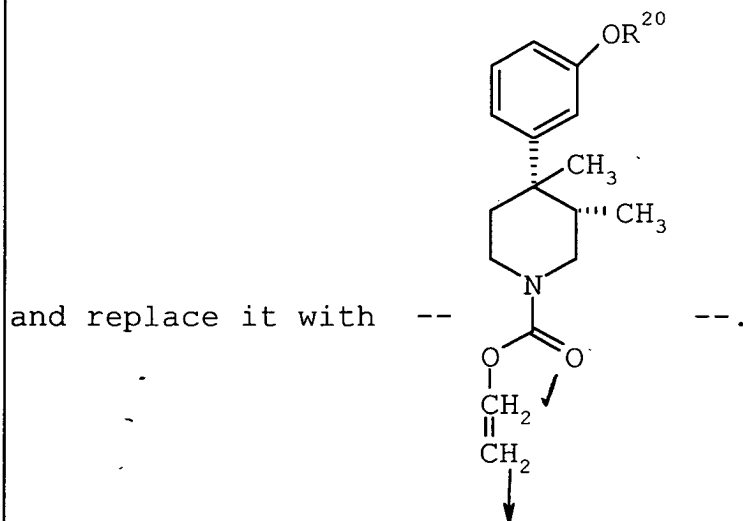
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It is certified that an error appears in the above-identified patent and that said Letters Patent is hereby corrected as shown below:



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✓

Column 17, line 27, delete the term "1,3,%-trimethyl" and replace it with --1,3,4-trimethyl--.

Column 17, line 48, delete the term "[ $\alpha$ ]<sub>589</sub>" and replace it with --[ $\alpha$ ]<sub>365</sub>--.

Column 22, line 47, delete the term "C<sub>26</sub>H<sub>35</sub>N<sub>3</sub>O·HCl" and replace it with --C<sub>26</sub>H<sub>35</sub>N<sub>3</sub>O<sub>3</sub>·HCl--.

Column 22, line 63, delete the term "424<sup>+/+</sup>" and replace it with --424 M<sup>+/+</sup>--.

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DATED : October 5, 1993

INVENTOR(S) : Buddy E. Cantrell, et al.

It is certified that an error appears in the above-identified patent and that said Letters Patent is hereby corrected as shown below:

Column 23, line 7, delete the term "U-NHCH<sub>3</sub>C(O)-NHCH<sub>2</sub>CH<sub>3</sub>·HCl·H<sub>2</sub>O]" and replace it with --[U-NHCH<sub>2</sub>C(O)-NHCH<sub>2</sub>CH<sub>3</sub>·HCl·H<sub>2</sub>O]--.

Column 23, line 53, delete the term "[G-NH(CH<sub>3</sub>)<sub>2</sub>C(O)NH<sub>2</sub>·HCl]" and replace it with --[G-NH(CH<sub>2</sub>)<sub>2</sub>C(O)NH<sub>2</sub>·HCl].

Column 25, line 21, delete the term "119°-12° C" and replace it with --119°-124°C--.

Column 27, line 9, delete the term "NHCH<sub>2</sub>C(O)OCH<sub>2</sub>CH<sub>2</sub>·HCl]" and replace it with --NHCH<sub>2</sub>C(O)OCH<sub>2</sub>CH<sub>3</sub>·HCl]--.

Column 28, line 67, delete the term "65.0S" and replace it with --65.05--.

Column 29, line 58, delete the term "C, 6%.91" and replace it with --C, 64.91--.

Column 29, line 59, delete the term "C, 6S.04" and replace it with --65.04--.

Column 30, line 17, delete the term "6S.86" and replace it with --65.86--.

Column 30, line 53, delete the term "444 M+1" and replace it with --444 M<sup>+</sup>+1--.

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It is certified that an error appears in the above-identified patent and that said Letters Patent is hereby corrected as shown below:

Column 35, line 10, delete the term "[X-O-(CH<sub>6</sub>H<sub>11</sub>)·HCl]" and replace it with --"[X-O-(C<sub>6</sub>H<sub>11</sub>)·HCl]".

Column 35, line 34, delete the term "3,%-dimethyl" and replace it with --3,4-dimethyl--.

Column 35, line 36, delete the term "[X-OCH<sub>2</sub>(CH<sub>6</sub>H<sub>11</sub>)·HCl]" and replace it with --[X-OCH<sub>2</sub>(C<sub>6</sub>H<sub>11</sub>)·HCl]--.

Column 35, line 39, delete the term "(7S0 mg)" and replace it with --(750 mg)--.

Column 36, line 8, delete the term "C<sub>298</sub>" and replace it with --C<sub>29</sub>--.

Column 36, line 17 delete the term "[XOCH<sub>2</sub>(CH<sub>6</sub>H<sub>5</sub>)·HCl]" and replace it with --[XOCH<sub>2</sub>(C<sub>6</sub>H<sub>5</sub>)·HCl]--.

Column 36, line 45, delete the term "[Z-NH(CH<sub>2</sub>)<sub>2</sub>C(O)OCH<sub>2</sub>(CH<sub>6</sub>H<sub>5</sub>)·HCl]" and replace it with --[Z-NH(CH<sub>2</sub>)<sub>2</sub>C(O)OCH<sub>2</sub>(C<sub>6</sub>H<sub>5</sub>)·HCl]--.

Column 37, line 9, delete the term "Hz" and replace it with --H<sub>2</sub>--.

Column 38, line 47, delete the term "C<sub>26</sub>" and replace it with --C<sub>29</sub>--.

Column 39, line 1, after the term "HCl" add -----.

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It is certified that an error appears in the above-identified patent and that said Letters Patent is hereby corrected as shown below:

Column 39, line 9, after the term "monohydrochloride" add --. [Z--.

Column 39, line 21, delete the term "C, 66.4" and replace it with --66.44--.

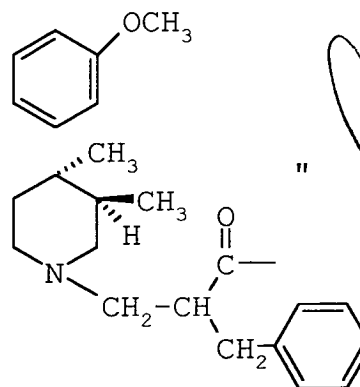
Column 40, line 49, delete the term "C, 63 77" and replace it with --C, 63.77--.

Column 42, line 28, delete the term "3,4-dimethyl1-" and replace it with --3,4-dimethyl-1- --.

Column 42, line 60, delete the phrase "[H<sub>2</sub>NCH<sub>2</sub>C(O)NHCH<sub>2</sub>C(O)NHCH<sub>2</sub>C(O)NHCH<sub>2</sub>-(C<sub>6</sub>H<sub>5</sub>)]" and replace it with --[H<sub>2</sub>NCH<sub>2</sub>C(O)NHCH<sub>2</sub>C(O)NHCH<sub>2</sub>-(C<sub>6</sub>H<sub>5</sub>)]--.

Column 42, line 64, delete the term "t-butoxycarbon-yl" and replace it with --t-butoxycarbonyl--.

Column 45, lines 25-40, delete the structure "



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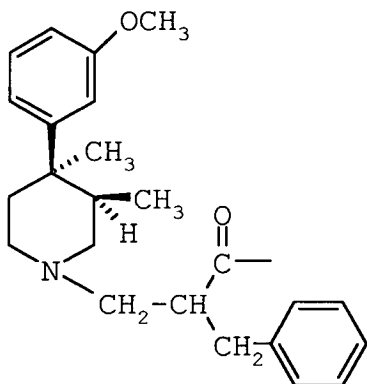
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and replace it with --



Column 45, line 65, delete the term "(2..72 g)" and replace it with --2.72 g)--.

Column 49, line 52, delete the term "218 m9" and replace it with --218 mg--.

Column 49, line 66, delete the term "aceate" and replace it with --acetate--.

Column 51, line 20, delete the term "H, 7.0B;" and replace it with --H, .7.08;--.

Column 51, line 33, delete the term "N<sub>3</sub>OH" and replace it with --NaOH--.

Column 53, line 11, delete the term "N<sub>3</sub>OH" and replace it with --NaOH--.

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DATED : October 5, 1993

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Column 53, line 40, delete the term "57-67" and replace it with --57-65--

Column 53, line 66, delete the term "ms (fd)'424 M<sup>+</sup>," and replace it with --ms (fd) = 424 M<sup>+</sup>--.

Column 55, line 7, delete the term "7.7N 6.32;" and replace it with --7.74 N 6.32;--.

Column 57, line 23, after the graph, insert the following

- (1) compound tested corresponds to Example Number
- (2) mg/kg in mouse writhing test
- (3) mg/kg in mouse diarrhea test
- (4) ration of AD<sub>50</sub> to ED<sub>50</sub>

\* Diastereomer A

\*\* Diastereomer B

\*\*\* I. V. administration because of lack of sample--.

Column 64, line 1, delete the term "0-;" and replace it with --0-4;--.

Column 64, line 46, delete the term "NHCH<sub>3</sub>C(O)NHCH<sub>2</sub>CH<sub>3</sub>," and replace it with -- NHCH<sub>2</sub>C(O)NHCH<sub>2</sub>CH<sub>3</sub>--.

Column 64, line 59, delete the term "(CH<sub>2</sub>C(O)OH," and replace it with --(CH<sub>2</sub>)<sub>2</sub>C(O)OH,--.

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Column 66, line 22, delete the term "ZNCH<sub>2</sub>C(O)OCH<sub>2</sub>CH(CH<sub>3</sub>)<sub>2</sub>," and replace it with --ZNHCH<sub>2</sub>C(O)OCH<sub>2</sub>CH(CH<sub>3</sub>)<sub>2</sub>--.

Column 66, line 32, delete the term "(C<sub>2</sub>)<sub>q</sub>-B," and replace it with --(CH<sub>2</sub>)<sub>q</sub>-B,--.

Column 66, line 45, delete the term "ZNCH<sub>2</sub>C(O)OCH<sub>2</sub>CH(CH<sub>3</sub>)<sub>2</sub>," and replace it with --ZNHCH<sub>2</sub>C(O)OCH<sub>2</sub>CH(CH<sub>3</sub>)<sub>2</sub>--.

Column 67, line 1, delete the term "ZNCH<sub>2</sub>C(O)OCH<sub>2</sub>CH(CH<sub>3</sub>)<sub>2</sub>," and replace it with --ZNHCH<sub>2</sub>C(O)OCH<sub>2</sub>CH(CH<sub>3</sub>)<sub>2</sub>--.

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